PTO/SB/21 (08-03) Approved for use through 07/31/2006. OMB 0651-0031

U.S. Patent and Trademark Office: U.S. DEPARTMENT OF COMMERCE

<i>f</i>		Application Number	10/789,222	
TRANSMITTAL		Filing Date	February 24, 2004	
FORM		First Named Inventor	Qin Yu	
(to be used for all correspondence after in	itial filing)	Art Unit	Not Yet Assigned	
		Examiner Name	Not Yet Assigned	
Total Number of Pages in This Submission	n 14	Attorney Docket Number	UPN0003-100 (P3115)	$\overline{\mathcal{J}}$
	ENCL	OSURES (check all that apply)		
Fee Transmittal Form	☐ Draw	ing(s)	After Allowance Communication to Group	
Fee Attached	Licen	sing-related Papers	Appeal Communication to Board of Appeals and Interferences	
Amendment / Reply	Petiti	on	Appeal Communication to Group (Appeal Notice, Brief, Reply Brief)	
After Final		on to Convert to a sional Application	Proprietary Information	
Affidavits/declaration(s)		er of Attorney, Revocation ge of Correspondence Address	Status Letter	
Extension of Time Request	☐ Term	inal Disclaimer	Other Enclosure(s) (please identify below):	
	Requ	est for Refund	References AA-DT	
Express Abandonment Request	☐ CD, I	Number of CD(s)		
☐ Information Disclosure Statement			,	

Remarks Document(s) Response to Missing Parts/ Incomplete Application Response to Missing Parts under 37 CFR 1.52 or 1.53 SIGNATURE OF APPLICANT, ATTORNEY, OR AGENT Firm David A. Sadewasser/Reg. No. 55,587 Individual name Signature Date

CERTIFICATE OF MAILING

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David A. Sadewasser

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IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In Re Application of: Qin Yu

Serial No.: 10/789,222 Group Art Unit: Not Yet Assigned

Filing Date: February 27, 2004 Examiner: Not Yet Assigned

For: ANGIOPOIETIN AND FRAGMENTS MUTANTS AND ANALOGS THEREOF AND

USES OF THE SAME

DATE OF DEPOSIT: May 24, 2604
I HEREBY CERTIFY THAT THIS PAPER IS BEING
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TYPED NAME: Dayld A. Sadewasser REGISTRATION NO:55,587

Commissioner for Patents P.O. Box 1450 Alexandria, VA 22313-1450

Dear Sir:

INFORMATION DISCLOSURE STATEMENT

Pursuant to 37 C.F.R. §§ 1.56 and in accordance with 37 C.F.R. §§ 1.97 and 1.98, information relating to the above-identified application is hereby disclosed, the Examiner in charge of the above-identified application is requested to consider and make of record the references listed on the PTO Form SB/08A and PTO Form SB/08B, formerly known as PTO Form 1449 submitted herewith.

Inclusion of the information submitted herewith is not to be construed as an admission that the information is material as that term is defined in 37 C.F.R. § 1.56(b).

In accordance with 37 C.F.R. § 1.97(g), the filing of this Information Disclosure Statement shall not be construed to mean that a search has been made.

This l	Inform	ation Disclosure Statement is being filed:
\boxtimes	withir	three months of the filing date of the patent application.
		three months of the date of entry into the national stage as set forth in F.R. § 1.491 of the international application.
	befor	e the mailing date of a first Office Action on the merits.
	mailir	the mailing date of a first Office Action on the merits, but before the ng date of a Final Office Action under 37 C.F.R. § 1.116 or a Notice of vance under 37 C.F.R. § 1.311, and accordingly is accompanied by:
		the Statement under 37 C.F.R. § 1.97(e) (see "Statement" below);
		or
		the Fee of \$180.00 set forth in 37 C.F.R. § 1.17(p); or
		No fee is owed by the applicant(s).
	Stater Final C.F.R	cordance with 37 C.F.R. § 1.129(a), this Information Disclosure ment is being filed in connection with \square the first or \square second After Submission, and accordingly is accompanied by the Statement under 37 \square § 1.97(e) (see "Statement" below) and the fee of \$180.00 as set forth in F.R. § 1.17(p), is attached.
	Notice with, Stater Inform	the mailing date of a Final Office Action under 37 C.F.R. § 1.116 or a e of Allowance under 37 C.F.R. § 1.311, but before, or simultaneously the payment of the Issue Fee, and accordingly is accompanied by the nent under 37 C.F.R. § 1.97(e), a Petition requesting consideration of the nation Disclosure Statement and the Petition Fee of \$130.00 set forth in F.R. § 1.17(i)(1) (see "Statement," "Petition," and "Fees" below).
		es of references (AA-DT) listed on the attached PTO Form SB/08A and Form SB/08B, formerly known as PTO Form 1449 are enclosed.
	EXC	EPT THAT:
		In view of the voluminous nature of reference @@, and the likelihood that this reference is available to the Examiner, copies are not enclosed herewith.
		In accordance with 37 C.F.R. § 1.98(d), copies of the following references listed on the attached PTO Form SB/08A and PTO Form SB/08B, formerly known as PTO Form 1449 are not enclosed herewith because they were previously cited by or submitted to the U.S. Patent and Trademark Office in patent application(s) for which a claim for priority under 35 U.S.C. § 120 have been made in the instant application.

	Copies of references listed on the attached PTO Form SB/08A a PTO Form SB/08B, formerly known as PTO Form 1449 we previously cited by or submitted to the U.S. Patent and Tradema Office in parent application Serial No. @@.	ere
	If any of the foregoing publications are not available to a Examiner, Applicant will endeavor to supply copies at a Examiner's request.	
Stater	ent under 37 C.F.R. § 1.97(e)	
	The undersigned attorney hereby states that each item information contain in the Information Disclosure Statement was cited in a communication from foreign patent office in a counterpart foreign patent application not more that three months prior to the filing of the Information Disclosure Statement.	n a
Stater	ent under 37 C.F.R. § 1.704(d)	
	The undersigned attorney hereby states that each item of information contained in the Information Disclosure Statement was cited in a communication from a foreign patent office in a counterpart application and the communication was not received by any individual designated in §1.56(c) more than 30 days prior to the filing of the Information Disclosure Statement.	
Fees		
\boxtimes	No Fee is owed by the applicant(s).	
	The Information Disclosure Statement Fee of \$180.00 under 37 C.F § 1.17(p) is enclosed herewith.	R.
	The Petition Fee of \$130.00 under 37 C.F.R. § 1.17(i)(1) is enclosed herewise	h.
Metho	d of Payment of Fees	
	Attached is a check in the amount of \$ This form submitted in duplicate.	is
	Charge Deposit Account No. 50-1275 in the amount of \$ This form is submitted in duplicate.	
\boxtimes	Please charge any deficiency or credit any overpayment to Deposit According 1275	ınt



No fee or Statement is required under 37 C.F.R. § 1.97(b).

Respectfully submitted,

Dated: May 24, 200

COZEN O'CONNOR, P.C. 1900 Market Street, 5th Floor Philadelphia, PA 19103-3508 (215) 665-2000 – Telephone (215) 701-2013 - Facsimile David A. Sadewasser Registration No. 55,587

PTO/SB/08b(08-03)

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R DE substitute for form 1449B/PTO

Sheet

INFORMATION DISCLOSURE STATEMENT BY APPLICANT

of

Complete if Known Application Number 10/789,222 February 27, 2004 Filing Date First Named Inventor Qin Yu Not Yet Assigned Art Unit Not Yet Assigned Examiner Name Attorney Docket Number UPN0003-100 (P3115)

(Use as many sheets as necessary) 9

NON PATENT LITERATURE DOCUMENTS Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of T² Cite Examiner the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue No. number(s), publisher, city and/or country where published. Initials * FOLKMAN, "Tumor angiogenesis: therapeutic implications," New. Eng. J. Med. (1971) 285:1182-1186 AA RISAU, "Mechanisms of angiogenesis," Nature (1997) 386:671-674. AB KIM, et al., "Inhibition of vascular endothelial growth factor-induced angiogenesis suppresses tumor growth in AC vivo," Nature (1993) 362:841-844. HANAHAN and FOLKMAN, "Patterns and emerging mechanisms fo the angiogenic switch during tumorigenesis," ΑD Cell (1996) 86:353-364. HANAHAN, "Signalling vascular morhogenesis and maintenance," Science (1997) 277:48-50. ΑE HANAHAN and WEINBERG, "The hallmarks of cancer," Cell (2000) 100:57-70. AF FOLKMAN and D'AMORE, "Blood vessel formation: what is its molecular basis?", Cell (1996) 87:1153-1155. AG YANCOPOULOS, et al., "Vascular-specific growth factors and blood vessel formation," Nature (2000) 407:242-INGBER and FOLKMAN, "How does extracellular matrix control capillary morphogenesis?", Cell (1989) 58:803-ΑI RAMSAUER and D'AMORE, "Getting tie(2)d up in angiogenesis," J. Clin. Investig. (2002) 110:1615-1617. AJ BETSHOLTZ, et al., "Developmental roles of platelet -derived growth factors," BioEssays (2001) 23:494-507. ΑK

Examiner	Date	
Signature	Considered	

^{*}EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

Applicant's unique citation designation number (optional). Applicant is to place a check mark here if English language Translation is attached.

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Substitute	e for form 1449B/PTO)		Complete if Known			
		.		Application Number	10/789,222		
			CLOSURE	Filing Date	February 27, 2004		
STAT	LEWENT B	Y A	PPLICANT	First Named Inventor	Qin Yu		
				Art Unit	Not Yet Assigned		
	(Use as many she	ets as	necessary)	Examiner Name	Not Yet Assigned	•	
Sheet	2	of	9	Attorney Docket Number	UPN0003-100 (P3115)		

		NON PATENT LITERATURE DOCUMENTS	,
Examiner Initials *	Cite No. ¹	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.	T²
	AL	FONG, et al., "Role of the Flt-1 receptor tyrosine kinase in regulating the assembly of vascular endothelium," Nature (1995) 376:66-70.	
	AM	MAISONPIERRE, et al., "Angiopoietin-2, a natural antagonist for tie2 that disrupts in vivo angiogenesis," Science (1997) 277:55-60.	
·	AN	SATO, et al., "tie-1 and tie-2 define another class of putative receptor tyrosine kinase genes expressed in early embryonic vascular system," Proc. Natl. Acad. Sci. USA (1993) 90:9355-9358.	
	AO	SCHNURCH and RISAU, "Expression of the tie-2, a member of a novel family of receptor tyrosine kinases, in the endothelial cell lineage," Development (1993) 119:957-968.	
	AP	DUMONT, et al., "Dominant-negative and targeted null mutations in the endothelial receptor tyrosine kinase, tek, reveal a critical role in vasculogenesis of the embryo," Genes Dev. (1994) 8:1897-1909.	
	AQ	COOGAN, et al., "Expression of tie2/tek in breast tumor vasculature provides a new marker for evaluation of tumor angiogenesis," Br. J. Cancer (1998) 77:51-56.	
	AR	SATO, et al., "Distinct roles of the receptor tyrosine kinases tie-1 and tie-2 in blood vessel formation," Nature (1995) 376:70-74.	
	AS	SURI, et al., "Requisite role of angiopoietin-1, a ligand for the TIE2 receptor during embryonic angiogenesis," Cell (1996) 87:1171-1180.	
	AT	GALE and YANCOPOULOS, "Growth factors acting via endothelial cell-specific receptor tyrosine kinases: VEGFs, angiopoietins, and ephrins in vascular development," Genes Dev. (1999) 13:1055-1066.	
	AU	SURI, et al., "Increased vascularization in mice overexpressing angiopoietin-1," Science (1998) 282:468-471.	
	AV	THURSTON, et al., "Leakage-resistant blood vessels in mice transenically overexpressing angiopoietin-1," Science (1999) 286:2511-2514.	

Examiner	Date]
Signature	 Considered	

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Substitute	e for form 144	B/PTO		Complete if Known			
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			CLOSURE	Filing Date	February 27, 2004		
STA	TEMEN	T BY A	PPLICANT	First Named Inventor	Qin Yu		
				Art Unit	Not Yet Assigned		
	(Use as ma	ny sheets as	necessary)	Examiner Name	Not Yet Assigned		
Sheet	3	of	9	Attorney Docket Number	UPN0003-100 (P3115)		

		NON PATENT LITERATURE DOCUMENTS	
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	AW	THURSTON, et al., "Angiopoietin-1 protects the adult vasculature against plasma leakage," Nature Med. (2000) 6:460-463.	
	AX	STRATMANN, et al., "Cell type-specific expression of angiopoietin-1 and angiopoietin-2 suggests a role in glioblastoma angiogenesis," Am. J. Pathol. (1998) 153:1459-1466.	
	AY	WITZENBICHLER, et al., "Chemotactic properties of angiopoietin-1 and -2, ligands for the endothelial-specific receptor tyrosine kinase tie2.," J. Biol. Chem. (1998) 273:18514-18521.	
	AZ	CARLSON, et al., "Direct cell adhesion to the angiopoietins mediated by integrins," J. Biol. Chem. (2001) 276:26516-26525.	
	ВА	PAPAPETROPOULOS, et al., "Angiopoietin-1 inhibits endothelial cell apoptosis via the Akt/survivin pathway," J. Biol. Chem. (2000) 275:9102-9105.	
	ВВ	KIM, et al., "Angiopoietin-1 regulates endothelial cell survival through the phosphatidylinositol 3'-kinase/Akt signal transduction pathway," Circulation Res. (2000) 86:24-29.	
	вс	HAYES, et al., "Angiopoietin-1 and its receptor Tie-2 participate in the regulation of capillary-like tubulin formation and survival of endothelial cells," Microvasc. Res. (1999) 58:224-237.	
	BD	OH, et al., "Hypoxia and vascular endothelial growth factor selectively upregulate angiopoietin-2 in bovine microvascular endothelial cells," J. Biol. Chem. (1999) 274:15732-15739.	
	BE	MANDRIOTA and PEPPER, "Regulation of angiopoietin-2 mRNA levels in bovine microvascular endothelial cells by cytokines and hypoxia," Circulation Res. (1998) 83:852-859.	
	BF	KIM, et al., "Tumor necrosis factor-alpha upregulates angiopoietin-2 in human umbilical vein endothelial cells," Biochem. Biophys. Res. Comm. (2000) 269:361-365.	
	BG	KIM, et al., "Angiopoietin-1 induces endothelial cell sprouting through the activation of focal adhesion kinase and plasmin secretion," Circulation Res. (2000) 86:952-959.	

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Signature	Considered	

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Substitute	for form 1449B/PTO			Complete if Known		
	DILATION			Application Number	10/789,222	
	RMATION			Filing Date	February 27, 2004	
STAT	STATEMENT BY APPLICANT			First Named Inventor	Qin Yu	
				Art Unit	Not Yet Assigned	
	(Use as many shee	ets as ı	necessary)	Examiner Name	Not Yet Assigned	
Sheet	4	of	9	Attorney Docket Number	UPN0003-100 (P3115)	

		NON PATENT LITERATURE DOCUMENTS	
Examiner Initials *	Cite No.1	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.	T²
	вн	VALENZUELA, et al., "Angiopoietins 3 and 4: diverging gene counterparts in mice and humans," Proc. Natl. Acad. Sci. USA (1999) 96:1904-1909.	
	ВІ	SIEMEISTER, et al., "Two independent mechanisms essential for tumor angiogenesis: inhibition fo human melanoma xenograft growth by interfering with eiter the vascular endothelial growth factor receptor pathway of the tie-2 pathway," Cancer Res. (1999) 59:3185-3193.	
	BJ	MILLAUER, et al., "Glioblastoma growth inhibited in vivo by a dominant-negative Flk-1 mutant," Nature (1994) 367:576-579.	
	вк	GOLDMAN, et al., "Paracrine expression of a native soluble vascular endothelial growth factor receptor inhibits tumor growth, metastsis, and mortality rate," Proc. Natl. Acad. Sci. USA (1998) 95:8795-8800.	
	BL	AHMAD, et al., "The effects of angiopoietin-1 and -2 on tumor growth and angiogenesis in human colon cancer," Cancer Res. (2001) 61:1255-1259.	
	ВМ	ETOH, et al., "Angiopoietin-2 is related to tumor angiogenesis in gastric carcinoma:possible in vivo regulation via induction of proteases," Cancer Res. (2001) 61:2145-2153.	
	BN	HAWIGHORST, et al., "Activation of the tie2 receptor by angiopoietin-1 enhances tumor vessel maturation nad impairs squamous cell carcinoma growth," Am. J. Pathol. (2002) 100:1381-1392.	
	во	KOGA, et al., "Expression of angiopoletin-2 in human glioma cells and its role for angiogenesis," Cancer Res. (2001) 61:6248-6254.	
	ВР	PAPETTI and HERMAN, "Mechanisms of normal and tumor-derived angiogenesis," Am. J. Physiol. Cell Physiol. (2002) 282:C947-C970.	
	BQ	TEICHERT-KULISZEWSKA, et al., "Biological action of angiopoietin-2 in a fibrin matrix model of angiogenesis is associated with activation if Tie2," Cardiovasc. Res. (2001) 49:659-670.	

Examiner	Date	·
Signature	Considered	j

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Substitute for form 1449B/PTO				Complete if Known		
WIFO	DRAKTIC	NI DIO	OL OCUPE	Application Number	10/789,222	
			CLOSURE	Filing Date	February 27, 2004	
STAT	STATEMENT BY APPLICANT			First Named Inventor	Qin Yu	
				Art Unit	Not Yet Assigned	
(Use as many sheets as necessary)			necessary)	Examiner Name	Not Yet Assigned	
Sheet	5	of	9	Attorney Docket Number	UPN0003-100 (P3115)	

		NON PATENT LITERATURE DOCUMENTS		
Examiner Initials *	Cite No.1 Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.			
	BS	YU and STAMENKOVIC, "Localization of matrix metalloproteinase 9 to the cell surface provides a mechanism for CD44-mediated tumor invasion," Genes Dev. (1999) 13:35-48.		
	ВТ	HUNGERFORD and LITTLE, "Developmental biology of the vascular smooth muscle cell: building a multilayed vessel wall," J. Vasc. Res. (1999) 36:2-27.		
	BU	GALE, et al., "Angiopoietin-2 is required for postnatal angiogenesis and lymphatic patterning, and only the latter role is rescued by angiopoietin-1," Devel. Cell (2002) 3:411-423.		
	BV	SHYU, et al., "Direct intramuscular injection of plasmid DNA encoding angiopoietin-1 but not angiopoietin-2 augments revascularization in the rabbit ischemic hindlimb," Circulation (1998) 98:2081-2087.		
	вw	KIM, et al., "Angiopoietin-2 at high concentration can enhance endothelial cell survival through the phosphatidylinositol 3'-kinase/Akt signal transduction pathway," Oncogene (2000) 19:4549-4552.		
	вх	LANDER and SELLECK, "The elusive functions of proteoglycans: in vivo veritas," J. Cell Biol. (2000) 148:227-232.		
	BY	IOZZO, "Matrix metalloproteins: from molecular design to cellular function," Ann. Rev. Biochem. (1998) 67:609-652.		
	BZ	IOZZO and SAN ANTONIO, "Heparan sulfate proteoglycans: heavy hitters in the angiogenesis arena," J. Clin. Investig. (2001) 108:349-355.		
	CA	FIEDLER, et al., "Angiopoietin-1 and angiopoietin-2 share the same binding domains in the tie-2 receptor involving the first Ig-like loop and the epidermal growth factor-like repeats," J. Biol. Chem. (2003) 278:1721-1727.		
	СВ	YU, et al., "Induction of apoptosis of metastatic mammary carcinoma cells in vivo by disruption of tumor cell surface CD44 function," J. Exp. Med. (1997) 186:1985-1996.		
	сс	KONTOS, et al., "Tyrosine 1011 of tie2 is the major site of association of p85 and is required for activation of phosphatidylinositol 3-kinase and Akt," Mol. Cell. Biol. (1998) 18:4131-4140.		

Examiner	Date	
Signature	Considered	

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	D114710N	DIO	OL COLLDE	Application Number	10/789,222
			CLOSURE	Filing Date	February 27, 2004
STAT	STATEMENT BY APPLICANT			First Named Inventor	Qin Yu
				Art Unit	Not Yet Assigned
(Use as many sheets as necessary)			necessary)	Examiner Name	Not Yet Assigned
Sheet	6	of	9	Attorney Docket Number	UPN0003-100 (P3115)

		NON PATENT LITERATURE DOCUMENTS	,
Examiner Initials *	Cite No.1	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.	T ²
	CD	FIDLER and ELLIS, "The implications of angiogenesis for the biology and therapy of cancer metastasis," Cell (1994) 79:185-188.	
_	CE	FIDLER, "Angiogenetic heterogeneity: regulation of neoplastic angiogenesis by the organ microenvironment," J. Natl. Cancer Inst. (2001) 93:1040-1041.	
	CF	ALI, et al., "Estrogen receptor-alpha in the inhibition of cancer growth and angiogenesis," Cancer Res. (2000) 60:7094-7098.	
	CG	NOKIHARA, et al., "Natural killer cell-dependent suppression of systemic spread of human lung adenocarcinoma cells by monocyte chemoattractant protein-1 gene transfection in severe combined immunodeficient mice," Cancer Res. (2000) 60:7002-7007.	
	СН	LINDAHL, et al., "Pericyte loss and microaneurysm formation in PDGF-B-deficient mice," Science (1997) 277:242-245.	
	СІ	GENGRINOVITCH, et al., "Glypican-1 is a VEGF165 binding proteoglycan that acts as an extracellular chaperone for VEGF165," J. Biol. Chem. (1999) 274:10816-10822.	
	C1	LI, et al., "Increased responsiveness of hypoxic endothelial cells to FGF2 is mediated by HIF-1alpha-dependent regulation of enzymes involved in synthesis of heparan sulfate FGF2-binding sites," J. Cell Sci. (2002) 115:1951-1959.	
	СК	NEUFELD, et al., "Vascular endothelial growth factor (VEGF) and its receptors," FASEB J. (1999) 13:9-22.	_
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	CN	XU and YU, "E-cadherin negatively regulates CD44-hyaluronan interaction and CD44-mediated tumor invasion and branching morphogenesis," J. Biol. Chem. (2003) 278:8661-8668.	

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INFORMATION DISCLOSURE				Filing Date	February 27, 2004	
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				Examiner Name	Not Yet Assigned	
Sheet	7	of	9	Attorney Docket Number	UPN0003-100 (P3115)	1

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	со	POLTORAK, et al., "VEGF145, a secreted vascular endothelial growth factor isoform that binds to extracellular matrix," J. Biol. Chem. (1997) 272:7151-7158.	
	СР	ROBINSON and STRINGER, "The splice variants of vascular endothelial growth factor (VEGF) and their receptors," J. Cell Sci. (2001) 114:853-865.	
	ca	RUHRBERG, "Endogenous inhibitors of angiogenesis," J. Cell Sci. (2001) 114:3215-3216.	
	CR	SAARISTO, et al., "Mechanisms of angiogenesis and their use in the inhibition of tumor growth and metastasis," Oncogene (2000) 19:6122-6129.	
	cs	MAESHIMA, et al., "Tumstatin, an endothelial cell-specific inhibitor of protein synthesis," Science (2002) 295:140-143.	
	СТ	O'REILLY, et al., "Angiostatin: a novel angiogenesis inhibitor that mediates the suppression of metastases by a Lewis lung carcinoma," Cell (1994) 79:315-328.	
	CU	O'REILLY, et al., "Antiangiogenic activity of the cleaved conformation of the serpin antithrombin," Science (1999) 285:1926-1928.	
	cv	YI and RUOSLAHTI, "A fibronectin fragment inhibits tumor growth, angiogenesis, and metastasis," Proc. Natl. Acad. Sci. USA (2001) 98:620-624.	-
	cw	VU, et al., "MMP-9/gelatinase-B is a key regulator of growth plate angiogenesis and apoptosis of hypertrophic chondrocytes," Cell (1998) 93:411-422.	
	сх	VAJKOCZY, et al., "Microtumor growth initiates angiogenic sprouting with angiogenic sprouting with simultaneous expression of VEGF, VEGF receptor-2, and angiopoietin-2," J. Clin. Investig. (2002) 109:777-785.	
	CY	BLOEMENDAL, et al., "New strategies in anti-vascular cancer therapy," Eur. J. Clin. Investig. (1999) 29:802-809.	ļ

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	cz	HARFOUCHE, et al., "Mechanisms which mediate the antiapoptotic effects of angiopoietin-1 on endothelial cells," Microvasc. Res. (2002) 64:135-147.	
	DA	HIRAOKA, et al., "Matrix metalloproteinases regulate neovascularization by acting as pericellular fibrinolysins," Cell (1998) 95:365-377.	
	DB	BERGERS, et al., "Matrix metalloproteinase-9 triggers the angiogenic switch during carcinogenesis," Nature Cell Biol. (2000) 2:737-744.	
	DC	FANG, et al., "Matrix metalloproteinase-2 is required for the switch to the angiogenic phenotype in a tumor model," Proc. Natl. Acad. Sci. USA (2000) 97:3884-3889.	
	DD	PFEIFER, et al., "Suppression of angiogenesis by lentiviral delivery of PEX, a noncatalytic fragment of matrix metalloproteinase 2," Proc. Natl. Acad. Sci. USA (2000) 97:12227-12232.	
	DE	STERNLICHT and WERB, "How matrix metalloproteinases regulate cell behavior," Ann. Rev. Cell Dev. Biol. (2001) 17:463-516.	
	DF	SILLETTI, et al., "Disruption of matrix metalloproteinase 2 binding to integrin alphavbeta3 by an organic molecule inhibits angiogenesis and tumor growth in vivo," Proc. Natl. Acad. Sci. USA (2001) 98:119-124.	
	DG	SIPES, et al., "Cooperation between thrombospondin-1 type 1 repeat peptides and alphavbeta3 integrin ligands to promote melanoma cell spreading and focal adhesion kinase phosphorylation," J. Biol. Chem. (1999) 274:22755-22762.	
	DH	VISCONTI, et al., "Orchestration of angiogenesis and arteriovenous contribution by angiopoietins and vascular endothelial groth factor (VEGF)," Proc. Natl. Acad. Sci. USA (2002) 99:8219-8224.	
	DI	UEMURA, et al., "Recombinant angiopoietin-1 restores higher-order architecture of growing blood vessels in mice in the absence of mural cells," J. Clin. Invest. (2002) 110:1619-1628.	
	DJ	YU and STMENKOVIC, "Cell surface-localized matrix metalloproteinase-9 protelytically activates TGF-beta and promotes tumor invasion and angiogenesis," Genes Dev. (2000) 14:163-176.	

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	DK	McFALL and RAPRAEGER, "Characterization of the high affinity cell-binding domain in the cell surface proteoglycan syndecan-4," J. Biol. Chem. (1998) 273:28270-28276.	
	DL	OLSON, et al., "High affinity binding of latent matrix metalloproteinase-9 to the alpha2(IV) chain of collagen IV," J. Biol. Chem. (1998) 273:10672-10681.	
	DM	BROOKS, et al., "Localization of matrix metalloproteinase MMP-2 to the surface of invasive sells by interaction with integrin alphavbeta3," Cell (1996) 85:683-693.	
	DN	MOYON, et al., "Selective expression of angiopoietin 1 and 2 in mesenchymal cells surrounding veins and arteries of the avian embryo," Mechs. Devel. (2001) 106:133-136.	
	DO	WONG, et al., "Tie2 expression and phosphorylation in angiogenic and quiescent adult tissues," Circ. Res. (1997) 81:567-574.	
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	DR	JOUSSEN, et al., "Suppression of diabetic retinopathy with angiopoietin-1," Am. J. Pathol. (2002) 160:1683-1693.	
	DS	HATTORI, et al., "Vascular endothelial growth factor and angiopoietin-1 stimulate postnatal hematopoiesis by recruitment of vasculogenic and hematopoietic stem cells," J. Exp. Med. (2001) 193:1005-1014.	
	DT	DAVIS, et al., "Angiopoietins have distinct modular domains essential for receptor binding, dimerization and superclustering," Nature Struct. Biol. (2002) 10:38-44.	

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